# (19) World Intellectual Property Organization

International Bureau





#### (43) International Publication Date 16 February 2006 (16.02.2006)

(10) International Publication Number WO 2006/015490 A1

(51) International Patent Classification<sup>7</sup>: 27/54, A61F 2/14, G02B 1/00

A61L 27/24,

(21) International Application Number:

PCT/CA2005/001240

**(22) International Filing Date:** 12 August 2005 (12.08.2005)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

60/601,270

13 August 2004 (13.08.2004) U

- (71) Applicants (for all designated States except US): OTTAWA HEALTH RESEARCH INSTITUTE [CA/CA]; 725 Parkdale Avenue, Ottawa, Ontario K1Y 4E9 (CA). NATIONAL RESEARCH COUNCIL OF CANADA [CA/CA]; 1200 Montreal Road, Bldg. M-58, Room EG12, Ottawa, Ontario K1A 0R6 (CA).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): GRIFFITH, May [CA/CA]; R.R.#3, 1006 Spruce Ridge Road, Carp, Ontario K0A 1L0 (CA). CARLSSON, David, J. [CA/CA]; 14 Delong Drive, Ottawa, Ontario K1J 7E6 (CA). LI, Fengfu [CA/CA]; 19 Maxton Private, Ottawa, Ontario K1J 1J1 (CA). LIU, Yuwen [CA/CA]; 514--790 Springfield Drive, Ottawa, Ontario K1V 6L2 (CA). ASMANRAFAT, Merhdad [CA/CA]; 112 Robert-Stewart Road, Gatineau, Québec J9H 7K7 (CA).

- (74) Agents: WHITE, Stephanie, R. et al.; Osler, Hoskin & Harcourt LLP, Suite 1500, 50 O'Connor Street, Ottawa, Ontario K1P 6L2 (CA).
- (81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW.
- (84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European (AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

#### Published:

- with international search report
- before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

#### (54) Title: OPHTHALMIC DEVICES AND RELATED METHODS AND COMPOSITIONS

(57) Abstract: Devices, methods, and compositions for improving vision or treating diseases, disorders or injury of the eye are described. Ophthalmic devices, such as corneal onlays, corneal inlays, and full-thickness corneal implants, are made of a material that is effective in facilitating nerve growth through or over the device. The material may include an amount of collagen greater than 1% (w/w), such as between about 10% (w/w) and about 30% (w/w). The material may include collagen polymers and/or a second biopolymer or water-soluble synthetic polymer cross-linked using EDC/NHS chemistry. The material may additionally comprise a synthetic polymer. The devices are placed into an eye to correct or improve the vision of an individual or to treat a disease, disorder or injury of an eye of an individual.



ernational application No. PCT/CA2005/001240

### Box No. II Observations where certain claims were found unsearchable (Continuation of item 2 of the first sheet)

Thi reas			rnational search report has not been established in respect of certain claims under Article 17(2)(a) for the following
1.	[2	x]	Claim Nos.: 27
			because they relate to subject matter not required to be searched by this Authority, namely:
			claim 27 is directed to a method of treatment of the human or animal body (Rule 39.1 (iv) PCT), however, the search was carried out on the alleged effects of the composition.
2.	[	]	Claim Nos.:  because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
3.	Ε	]	Claim Nos.:  because they are dependant claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box	N	0.	Observations where unity of invention is lacking (Continuation of item 3 of first sheet)
1.	[	]	As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
2.	[	]	As all searchable claims could be searched without effort justifying additional fees, this Authority did not invite payment of additional fees.
3.	[	]	As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claim Nos. :
4.	[	]	No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claim Nos.:
			Remark on Protest [ ] The additional search fees were accompanied by the applicant's protest and, where applicable, the payment of a protest fee.
			[ ] The additional search fees were accompanied by the applicant's protest but the applicable protest fee was not paid within the time limit specified in the invitation.
			[ ] No protest accompanied the payment of additional search fees.

International application No. PCT/CA2005/001240

A. CLASSIFICATION OF SUBJECT MATTER IPC(7): A61L 27/24, A61L 27/54, A61F 2/14, G02B 1/00

According to International Patent Classification (IPC) or to both national classification and IPC

#### B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC(7): A61L-27, G02B-1, A61F-2/14

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic database(s) consulted during the international search (name of database(s) and, where practicable, search terms used) Canadian Patent Database, WEST (USPTO), PubMed

Keywords = collagen; cross\*link\*; corneal, eye, ophthalm\*; clear, transparent and translucent

#### C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 4,581,030 (Bruns, R. B. et al.) 8 April 1986 (08.04.1986) Abstract, Col. 2-3 and Examples 1 and 2	1-7, 13, 16, 18, 19, 23 and 27-29
Y	;	8-12, 17, 21, 22 and 26
X	US 4,223,984 (Miyata, T. et al.) 23 September 1980 (23.09.1980) Entire document	1-7, 13-16, 18, 19, 23, and 27-29
Y	1	8-12, 17, 21, 22 and 26
X	US 2003/0018123 (Bagrov, S. N. et al.) 23 January 2003 (23.01.2003) Entire document	1-6, 16, 19, 23 and 27-29
Y	<u>:</u>	8-12, 17, 21, 22 and 26
Y	Li, F. et al. "Cellular and nerve regeneration within a bioprosthetic extracellular matrix for corneal implantation" Proc. Nat. Acad. Sci. 2003 (Dec. 23), Vol. 100 (No. 26), pages 15346-15351.  Entire document	8-12, 17, 21, 22 and 26

[X]	Further documents are listed in the continuation of Box C.	[X]	See patent family annex.
*	Special categories of cited documents:	"T"	later document published after the international filing date or priority date and not in conflict with the application but cited to understand
"A"	document defining the general state of the art which is not considered to be of particular relevance		the principle or theory underlying the invention
"E"		"X"	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive
٦	earlier application or patent but published on or after the international filing date		step when the document is taken alone
"L"	document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"Y"	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art
"o"	document referring to an oral disclosure, use, exhibition or other means		
l ""		"&"	document member of the same patent family

"P" document published prior to the international filing date but later than the priority date claimed	"&" document member of the same patent family
Date of the actual completion of the international search	Date of mailing of the international search report
24 November 2005 (24-11-2005)	7 December 2005 (07-12-2005)
Name and mailing address of the ISA/CA Canadian Intellectual Property Office	Authorized officer
Place du Portage I, C114 - 1st Floor, Box PCT 50 Victoria Street	Stephen Decker (819) 934-2333
Gatineau, Quebec K1A 0C9 Facsimile No.: 001(819)953-2476	

International application No. PCT/CA2005/001240

ategory*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
P, Y	Li, F. et al. "Recruitment of multiple cell lines by collagen-synthetic copolymer matrices in corneal regeneration" Biomaterials 2005 (Jun.), Vol. 26 (No. 16), pages 3093-3104.  Entire document	8-12, 17, 21, 22 and 26
Y	Carlsson, D. J. et al. "Bioengineered corneas: how clsoe are we?" Curr. Opin. Ophthalmol. 2003 (Aug.), Vol. 14 (No. 4), pages 192-197. Pages 195-196	20
A	Leskul, M. et al. "CxGELSIX: a novel preparation of type VI collagen with possible use as a biomaterial" Cornea 2000 (Mar.), Vol. 19 (No. 2), pages 194-203.	1-29
A	Doillon, C. J. et al. "A collagen-based scaffold for a tissue engineered human cornea: physical and physiological properties" Int. J. Artif. Organs 2003 (Aug.), Vol. 26 (No. 8), pages 764-773.	1-29
A	CA 2,134,744 (Rhee, W. M. et al.) 4 May 1995 (04.05.1995)	1-29
A	CA 2,286,718 (Peyman, G. A.) 5 November 1998 (05.11.1998)	1-29

Information on patent family members

International application No. PCT/CA2005/001240

Patent Document Cited in Search Report	Publication Date	Patent Family Member(s)	Publication Date <sup>(</sup>	
US4581030	08-04-1986	US4505855	19-03-1985	
US4223984	23-09-1980	NONE		
US2003018123	23-01-2003	RU2190841 C1	10-10-2002	
CA2134744	04-05-1995	AT157351 T AT168708 T	15-09-1997 15-08-1998	
		AT197058 T AT255918 T	15-11-2000 15-12-2003	
		AU638687 B2 AU668265 B2	08-07-1993 26-04-1996	
		AU677789 B2 AU698233 B2	08-05-1997 29-10-1998	
		AU1507595 A AU4660989 A	09-11-1995 12-06-1990	
		AU4662093 A	31-01-1994	
		AU5522994 A CA2003538 A1	25-08-1994 21-05-1990	
		CA2115967 A1 CA2143923 A1	21-08-1994 03-11-1995	
		CA2172906 A1 DE59403818 D1	08-12-1996 02-10-1997	
		DE68928754 D1	27-08-1998	
		DE68928754 T2 DE69519124 D1	14-01-1999 23-11-2000	
		DE69519124 T2	17-05-2001	
		DE69630990 D1 DE69630990 T2	22-01-2004 30-12-2004	
		DK612723 T3 EP0444157 A1	30-03-1998 04-09-1991	
		EP0612723 A1	31-08-1994	
		EP0648239 A1 EP0656214 A1	19-04-1995 07-06-1995	
		EP0656215 A1 EP0680990 A1	07-06-1995 08-11-1995	
		EP0697218 A2	21-02-1996	
		EP0747066 A2 ES2107698 T3	11-12-1996 01-12-1997	
		ES2119743 T3 ES2211923 T3	16-10-1998 16-07-2004	
		FI114467 B1	29-10-2004	
		GR3024855 T3 HU70535 A2	30-01-1998 30-10-1995	
		IL108697 A	16-07-2000	
		JP2505312 B2 JP3554352 B2	05-06-1996 18-08-2004	
,		JP4502027 T JP7278203 A	09-04-1992 24-10-1995	
		JP7278312 A	24-10-1995	
		JP8053548 A JP8502082 T	27-02-1996 05-03-1996	
		JP9099052 A JP2004189755 A	15-04-1997 08-07-2004	
		NO300322 B1	12-05-1997	
		NZ250919 A TW420659 B	26-01-1996 01-02-2001	

Continued on Supplemental Page 6

International application No. PCT/CA2005/001240

Continued from Informa	tion on Patent Fam	nily Members (page 5)	
Patent Document Cited in Search Report	Publication Date	Patent Family Member(s)	Publication Date
		US5162430 US5264214 US5292802 US5304595 US5306500 US5308889 US5324775 US5328955 US5376375 US5413791 US5446091 US5470911 US5475052 US5476666 US5510121 US5510121 US5523348 US5527856 US5527856 US551087 US5550187 US5565519 US5614587 US5643464 US5744545 US5746421 US5800541 US5800541 US5800541 US58936035 WO9005755 A1	10-11-1992 23-11-1993 08-03-1994 19-04-1994 26-04-1994 03-05-1994 28-06-1994 12-07-1994 09-05-1995 29-08-1995 28-11-1995 12-12-1995 19-12-1995 19-12-1996 04-06-1996 04-06-1996 04-06-1996 15-10-1996 27-08-1996 25-03-1997 01-07-1997 28-04-1998 28-07-1998 01-09-1998 02-02-1999 10-08-1999 31-05-1990 20-01-1994 30-08-1994
CA2286718	05-11-1998	AU739297 B2 AU772492 B2 AU778310 B2 AU3713400 A AU3713500 A AU6456798 A BR0008601 A BR0008624 A BR9809289 A CA2364038 A1 CA2366022 A1 CN1253484 A CN1342095 A CN1360486 A EP1014872 A1 EP1158936 A1 EP1158933 A1 JP2000513986 T JP2002537895 A JP2002537895 A MXPA01008750 A	11-10-2001 29-04-2004 25-11-2004 21-09-2000 21-09-2000 24-11-1998 26-12-2001 26-12-2001 04-07-2000 08-09-2000 08-09-2000 17-05-2000 27-03-2002 24-07-2002 05-07-2000 05-12-2001 05-12-2001 05-12-2001 24-10-2000 12-11-2002 14-07-2003 14-07-2003
Continued on Supplement	al Page 7		

International application No. PCT/CA2005/001240

Continued from Suppler	nental Page 6		
Patent Document Cited in Search Report	Publication Date	Patent Family Member(s)	Publication Date
		US5722971 US5919185 US5964748 US6063073 US6197019 B1 US6203538 B1 US6217571 B1 US6221067 B1 US6280470 B1 US2001027314 A1 US2001034516 A1 WO0051526 A1 WO09848715 A1	03-03-1998 06-07-1999 12-10-1999 16-05-2000 06-03-2001 20-03-2001 17-04-2001 24-04-2001 28-08-2001 04-10-2001 25-10-2001 08-09-2000 08-09-2000 05-11-1998